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537736.trn
=> search combination or mbinator?
        167630 COMBINATION
                                                            08/537,736
           988 COMBINATOR?
        168549 COMBINATION OR COMBINATOR?
L1
=> s l1(2a)chemistry
        337636 CHEMISTRY
                                                                      SRMT
L2
            29 L1(2A) CHEMISTRY
=> d bib
     ANSWER 1 OF 29 MEDLINE
L2
                  MEDLINE
     96368010
AN
     Characterization of the complexity of small-molecule libraries of
TI
     electrospray ionization mass spectrometry.
     Dunayevskiy Y; Vouros P; Carell T; Wintner E A; Rebek J Jr
ΑU
     Department of Chemistry, Barnett Institute, Northeastern University,
CS
     Boston, Massachusetts 02115, USA.
     ANALYTICAL CHEMISTRY, (1995 Sep 1) 67 (17) 2906-15.
SO
     Journal code: 4NR. ISSN: 0003-2700.
     United States
CY
     Journal; Article; (JOURNAL ARTICLE)
DT
     English
LA
     9611
EM
=> d all
     ANSWER 1 OF 29 MEDLINE
L2
                  MEDLINE
     96368010
AN
     Characterization of the complexity of small-molecule libraries of
ΤI
     electrospray ionization mass spectrometry.
     Dunayevskiy Y; Vouros P; Carell T; Wintner E A; Rebek J Jr
ΑU
     Department of Chemistry, Barnett Institute, Northeastern University,
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     Boston, Massachusetts 02115, USA.
     ANALYTICAL CHEMISTRY, (1995 Sep 1) 67 (17) 2906-15.
SO
     Journal code: 4NR. ISSN: 0003-2700.
     United States
CY
     Journal; Article; (JOURNAL ARTICLE)
DT
LΑ
     English
     9611
EM
     The growing interest in ***combinatorial***
                                                        ***chemistry***
AΒ
     has led us to explore new analytical methods for the analysis of
     complex molecular libraries. Because an investigation of large
     mixtures with 10(4)-10(5) different chemical entities was not
      realistic, an alternative approach was pursued that included the
     analysis of small representative sublibraries using positive and
     negative ion electrospray mass spectrometry. The detailed analysis
     of these model mixtures, containing up to 55 components, allowed us
     to obtain important information about the composition of a library
     with considerable complexity. The results were used to improve the
      synthetic procedure in order to provide the maximum yield of
```

expected library components. The applicability of mass spectrometry

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to the analysis of complex matrices and the usefulness of the
      technique for scree g synthesized combinatori libraries to probe
      their expected diversity and complexity have been demonstrated.
 CT
       Amino Acid Sequence
      *Dipeptides: CH, chemistry
       Molecular Sequence Data
       Spectrum Analysis, Mass
CN
      0 (Dipeptides)
 => e review/dt
 E1
             231
                     RETRACTED PUBLICATION/DT
 E2
             198
                     RETRACTION OF PUBLICATION/DT
 E3
         517153 --> REVIEW/DT
 E4
          34742
                     REVIEW ACADEMIC/DT
         12627 REVIEW ACADEMIC/DT
12627 REVIEW LITERATURE/DT
4854 REVIEW MULTICASE/DT
25959 REVIEW OF REPORTED CASES/DT
268783 REVIEW TUTORIAL/DT
13 SCIENTIFIC INTEGRITY REVIEW/DT
 E5
 E6
E7
E8
E9
            392
250
E10
                     TECHNICAL REPORT/DT
E11
                     TWIN STUDY/DT
**** END OF FIELD ****
=> s e3-8
         517153 REVIEW/DT
          34742 "REVIEW ACADEMIC"/DT
          12627 "REVIEW LITERATURE"/DT
           4854 "REVIEW MULTICASE"/DT
          25959 "REVIEW OF REPORTED CASES"/DT
         268783 "REVIEW TUTORIAL"/DT
         517189 (REVIEW/DT OR "REVIEW ACADEMIC"/DT OR "REVIEW LITERATURE"/
L3
                 DT OR "REVIEW MULTICASE"/DT OR "REVIEW OF REPORTED CASES"/
                 DT OR "REVIEW TUTORIAL"/DT)
=> d his
      (FILE 'HOME' ENTERED AT 21:45:29 ON 10 OCT 96)
     FILE 'MEDLINE' ENTERED AT 21:45:32 ON 10 OCT 96
L1
          168549 SEARCH COMBINATION OR COMBINATOR?
L2
              29 S L1(2A) CHEMISTRY
                 E REVIEW/DT
L3
          517189 S E3-8
=> s 13 and 12
L4
              5 L3 AND L2
=> d 1-5 all
L4
     ANSWER 1 OF 5 MEDLINE
AN
     96166348
                 MEDLINE
     ***Combinatorial***
ΤI
                                  ***chemistry*** in the discovery and
     development of drugs.
```

```
ΑU
      Doyle P M
      Department of Medic l Chemistry, Wellcome Research Laboratories,
 CS
      Beckenham, Kent, UK.
      JOURNAL OF CHEMICAL TECHNOLOGY AND BIOTECHNOLOGY, (1995 Dec) 64 (4)
 SO
      317-24. Ref: 70
      Journal code: AL8. ISSN: 0268-2575.
 CY
      ENGLAND: United Kingdom
      Journal; Article; (JOURNAL ARTICLE)
 DT
        ***General Review; (REVIEW)***
        ***(REVIEW, TUTORIAL)***
 LA
      English
 FS
      Priority Journals; B
 EM
      9605
 CT
      Amino Acid Sequence
      Automation
      Databases, Factual
      *Drug Design
      Molecular Sequence Data
      Oligopeptides: CH, chemistry
      *Oligopeptides: CS, chemical synthesis
     0 (Oligopeptides)
CN
L4
     ANSWER 2 OF 5 MEDLINE
     96102839 MEDLINE
AN
     Solid-phase ***combinatorial***
TI
                                         ***chemistry*** and novel
     tagging methods for identifying leads.
     Chabala J C
ΑU
CS
     Phamacopeia Inc, Princeton, USA.
     CURRENT OPINION IN BIOTECHNOLOGY, (1995 Dec) 6 (6) 632-9. Ref: 24
SO
     Journal code: A92. ISSN: 0958-1669.
CY
     ENGLAND: United Kingdom
     Journal; Article; (JOURNAL ARTICLE)
DT
       ***General Review; (REVIEW)***
       ***(REVIEW, TUTORIAL)***
LA
     English
     Priority Journals
FS
EM
     9604
     Encoded combinatorial chemical synthesis on solid phase is a new
AΒ
     paradigm in organic chemistry that provides chemists with powers
     similar to those enjoyed by molecular biologists. Encoded chemical
     libraries will have a profound impact on all endeavors that seek to
     identify molecules with optimized properties and to understand the
     factors governing molecular interactions. In particular, the
     discovery and optimization of new therapeutic and diagnostic drug
     molecules, traditionally a slow manual process, will be greatly
    accelerated by this technology.
CT
      Amino Acid Sequence
     Biotechnology
      Chemistry, Organic
     Drug Design
     Molecular Sequence Data
     Oligonucleotides: GE, genetics
     Oligopeptides: CH, chemistry
```

\*Oligopeptides: CS, chemical synthesis Oligopeptides: GE, enetics 0 (Oligonucleotides); 0 (Oligopeptides) CNANSWER 3 OF 5 MEDLINE L4AN 96050146 MEDLINE TIStrategies and recent technologies in drug discovery. ΑU Kubinyi H Drug Design, BASF, Ludwigshafen. CS PHARMAZIE, (1995 Oct) 50 (10) 647-62. SO Ref: 311 Journal code: P4D. ISSN: 0031-7144. CYGERMANY: Germany, Federal Republic of Journal; Article; (JOURNAL ARTICLE) DT\*\*\*General Review; (REVIEW) \*\*\* \*\*\* (REVIEW, ACADEMIC) \*\*\* LAEnglish FS Priority Journals EM9603 In the last years, the paradigms of drug research changed AB significantly. New technologies were developed, in several different fields. \*\*\*Combinatorial\*\*\* \*\*\*chemistry\*\*\* and high-throughput screening increase our chances to find new lead structures, with less effort than by dedicated syntheses. Gene technology, in addition to providing therapeutically useful proteins, significantly contributes to rational drug design. The primary structure of a protein can be derived from the DNA sequence of the corresponding gene. Its relevance for a certain disease is investigated in transgenic animals. Expression of the protein in bacteria or in cell culture produces material for screening systems and for 3D structure determination by protein crystallography. NMR techniques, or electron cryo-microscopy. Structure-based and computer-aided design methods are applied to optimize lead structures with the least effort. A serious problem in the application of such techniques is their limitation to ligand-protein interactions. For the design of a therapeutically useful drug, also absorption, distribution, metabolism and elimination have to be considered. QSAR methods help in this respect. Scope and limitations of the new technologies are discussed in the context of conventional approaches in drug discovery. CTCheck Tags: Animal; Human; Support, Non-U.S. Gov't Genetic Engineering \*Pharmacology: TD, trends Research Design Technology, Pharmaceutical L4ANSWER 4 OF 5 MEDLINE AN 94176088 MEDLINE TI \*\*\*Combinatorial\*\*\* \*\*\*chemistry\*\*\* --applications of light-directed chemical synthesis. ΑU Jacobs J W; Fodor S P Affymax Research Institute, Palo Alto, CA 94304... CS TRENDS IN BIOTECHNOLOGY, (1994 Jan) 12 (1) 19-26. SO Ref: 29 Journal code: ALJ. ISSN: 0167-7799.

```
ENGLAND: United Kingdom
CY
     Journal; Article; ( JRNAL ARTICLE)
DT
       ***General Review; (REVIEW) ***
       ***(REVIEW, TUTORIAL) ***
     English
LΑ
FS
     В
     9406
EΜ
     Combinatorial methods in biology and chemistry are proving to be
AΒ
     powerful methods for generating molecular diversity. One approach,
     light-directed chemical synthesis, combines semiconductor-based
     photolithography technologies with solid-phase organic chemistry to
     synthesize large arrays of molecules with potential biological
     activity. This novel technology has the potential to provide
     libraries of both natural and synthetic molecules that might be
     screened rapidly for biological activity.
      Amino Acid Sequence
CT
      Base Sequence
      Biotechnology: MT, methods
      Carbamates: CH, chemistry
      Dynorphins: AA, analogs & derivatives
      Dynorphins: CH, chemistry
      Dynorphins: CS, chemical synthesis
      Dynorphins: GE, genetics
      DNA: GE, genetics
      Endorphins: CH, chemistry
      Endorphins: CS, chemical synthesis
      Endorphins: GE, genetics
      Molecular Sequence Data
      Oligonucleotides: CH, chemistry
     *Oligonucleotides: CS, chemical synthesis
      Oligonucleotides: GE, genetics
      Peptides: CH, chemistry
     *Peptides: CS, chemical synthesis
      Peptides: GE, genetics
     *Photochemistry: MT, methods
     74913-18-1 (Dynorphins); 83335-41-5 (rimorphin); 9007-49-2 (DNA)
RN
     0 (Carbamates); 0 (Endorphins); 0 (Oligonucleotides); 0 (Peptides)
CN
     ANSWER 5 OF 5 MEDLINE
L4
                  MEDLINE
     90058642
AN
     Polypeptide chain binding proteins: catalysts of protein folding and
TI
     related processes in cells.
     Rothman J E
ΑU
     Department of Biology, Princeton University, New Jersey 08544...
CS
     CELL, (1989 Nov 17) 59 (4) 591-601.
                                           Ref: 102
SO
     Journal code: CQ4. ISSN: 0092-8674.
CY
     United States
     Journal; Article; (JOURNAL ARTICLE)
DT
        ***General Review; (REVIEW) ***
        ***(REVIEW, ACADEMIC) ***
     English
LA
     Priority Journals; Cancer Journals
FS
EM
      9003
```

Subcellular compartments in which folding and assembly of proteins AB occur seem to have set of PCB proteins capabl of mediating these and related processes, such as translocation across membranes. When a domain of a polypeptide chain emerges from a ribosome during synthesis or from the distal side of a membrane during translocation, successive segments of the chain are incrementally exposed to solvent and yet are unlikely to be able to fold. This topological restriction on folding likely mandates a need for PCB proteins to prevent aggregation. Catalysis of topologically restricted folding by PCB proteins is likely to involve both an antifolding activity that postpones folding until entire domains are available and, more speculatively, a folding activity resulting from a programmed stepwise release that employs the energy of ATP hydrolysis to ensure a favorable pathway. We are left with a new set of problems. How do proteins fold in cells? What are the sequences or structural signals that dictate folding pathways? The new challenge will be to understand folding as a \*\*\*combination\*\*\* \*\*\*chemistry\*\*\* , enzymology, and cell biology. of physical

CT Check Tags: Animal

Antigens, Bacterial: ME, metabolism

\*Heat-Shock Proteins: ME, metabolism

\*Protein Conformation

CN 0 (Antigens, Bacterial); 0 (GroEL Protein); 0 (Heat-Shock Proteins)

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COST IN U.S. DOLLARS
SINCE FILE TOTAL
ENTRY SESSION
FULL ESTIMATED COST
2.80
2.95

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- L3 ANSWER 1 OF 1 SCISTRCH COPYRIGHT 1996 ISI (F
- AN 94:118410 SCISEARCH
- GA The Genuine Article (R) Number: MV090
- TI COMBINATORIAL CHEMISTRY APPLICATIONS OF LIGHT-DIRECTED CHEMICAL SYNTHESIS
- AU JACOBS J W (Reprint); FODOR S P A
- CS AFFYMAX RES INST, 4001 MIRANDA AVE, PALO ALTO, CA, 94304 (Reprint);
  AFFYMETRIX, SANTA CLARA, CA, 95051
- CYA USA
- SO TRENDS IN BIOTECHNOLOGY, (JAN 1994) Vol. 12, No. 1, pp. 19-26. ISSN: 0167-9430.
- DT General Review; Journal
- FS AGRI
- LA ENGLISH
- REC Reference Count: 29
- AB Combinatorial methods in biology and chemistry are proving to be powerful methods for generating molecular diversity. One approach, light-directed chemical synthesis, combines semiconductor-based photolithography technologies with solid-phase organic chemistry to synthesize large arrays of molecules with potential biological activity. This novel technology has the potential to provide libraries of both natural and synthetic molecules that might be screened rapidly for biological activity.
- CC BIOTECHNOLOGY & APPLIED MICROBIOLOGY
- STP KeyWords Plus (R): DRUG DISCOVERY; LIGANDS; LIBRARY; MOLECULES
- RF 92-0799 007; ANTIBODY ENGINEERING; ANTIGEN COMBINING SITE; FILAMENTOUS PHAGE; PROTEIN TARGETS 92-5823 001; B-CELL EPITOPES OF THE CHLAMYDIA-TRACHOMATIS MAJOR OUTER-MEMBRANE PROTEIN; PEPTIDE LIBRARIES; ANTIGENIC SITES; ANTIPEPTIDE ANTIBODIES

RE

Referenced Author (RAU)	Year  (RPY)	VOL (RVL)	PG  (RPG)	Referenced Work (RWK)
(IAO)		, (2002) +=====		 +==========
BARRETT R W	1985	6	113	NEUROPEPTIDES
BIRNBAUM S	1992	3	49	CURR OPIN BIOTECH
BRENNER S	1992	,   89	5381	P NATL ACAD SCI USA
BUNIN B A	1992	114	10997	J AM CHEM SOC
CARUTHERS M H	1985	230	281	SCIENCE
CHO C Y	1993	261	1303	SCIENCE
CWIRLA S E	1990	87	6378	P NATL ACAD SCI USA
DEVLIN J J	1990	249	404	SCIENCE
DOWER W J	1992	2	251	CURR BIOL
ELLINGTON A D	1990	346	818	NATURE
FODOR S P A	1993	364	555	NATURE
FODOR S P A	1991	251	767	SCIENCE
FRANK R	1992	48	9217	TETRAHEDRON
FURKA A	1991	37	487	INT J PEPT PROT RES
GEYSEN H M	1984	81	3998	P NATL ACAD SCI USA
HOLMES C P	1993		489	PERSPECTIVES MED CHE
HOUGHTEN R A	1991	354	84	NATURE

KERR J M	1993	115	2529	J AM CHEM_SOC
LAM K S	91	354	82	NATURE
MERRIFIELD R B	1963	85	2149	J AM CHEM SOC
MERRIFIELD R B	1986	232	341	SCIENCE
NEEDELS M C	1993	90	10700	P NATL ACAD SCI USA
NIKOLAIEV V	1993	6	161	PEPTIDE RES
PAVIA M R	1993	3	387	BIOORG MED CHEM LETT
PEASE A C				IN PRESS P NATL ACAD
SCOTT J K	1990	249	386	SCIENCE
SIMON R J	1992	89	9367	P NATL ACAD SCI USA
TUERK C	1990	249	505	SCIENCE
WELLS J A	1992	2	597	CURR OPIN STRUC BIOL